

STIC Search Report Biotech-Chem Library

To: Devesh Khare Location: REM-5C35&5C18

Art Unit: 1623

Tuesday, January 18, 2005

From: Beverly Shears Location: Remsen Bldg.

RM 1A54

Phone: 571-272-2528

beverly.shears@uspto.gov

Search Notes

Case Serial Numbers	10/058903	and	10/312951	attached.
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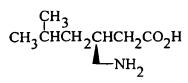


SEARCH REQUEST FORM

Scientific and Technical Information Center

	Chare Examiner #: 7 ber 272-0653 S ocation: 5C35 Results F	erial Number	: 10/058,903	_
If more than one search is submitt	ed, please prioritize	searches in	order of need.	Meg.
Please provide a detailed statement of the sear search include the elected species or structure: the concept or utility of the invention. Define citations, authors, etc, if known. Please attach	s, key words, synonyms, ac any terms that may have a	cronyms, and re special meaning	gistry numbers, and g. Give examples o	combine with
Title of Invention: See Bib Data She	eet on e-	•		
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Earliest priority Filing Date: 03/30/2 *For Sequence Searches Only* Please includ numbers) along with the appropriate serial numbers Please carry out a search on the serial numbers.	le all pertinent information umber.		divisional, or issued	l palent
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TO-1500 (9-00)



Pregabalin

Pregabalin Lactam

b-D-Lactose

b-Galactose

b-Glucose

Figure 1

HOOMUTO" WOOMMOON

APPIICVED	O F	IG.
by Dhaftsman	514	SUBCLASS 23

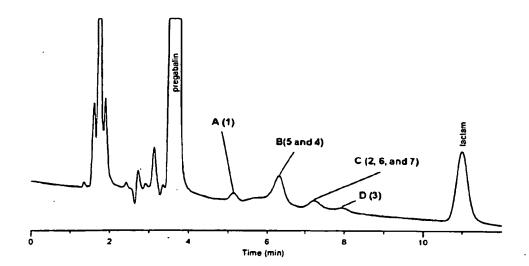


Figure 2. HPLC Chromatogram of Pregabalin 25-mg Capsule Stored at 40°C/75% RH for 6 Months (Original Method) HPLC Conditions: Column: Waters μ-Bondapak 10 μ C18, 300 × 3.9 mm ID; Mobile Phase: 550:350:100:1 H₂O:MeOH:CH₂ON:pH 7.0 Phosphate Buffer; Flow Rate: 1.0 mL/min; Detection: 210 nm; Temperature: Ambient; Compounds: as defined in Scheme 3

APPROVED	O1	FIG.
DHAFTSMAN	CLASS 514	SUBCLASS 2 2

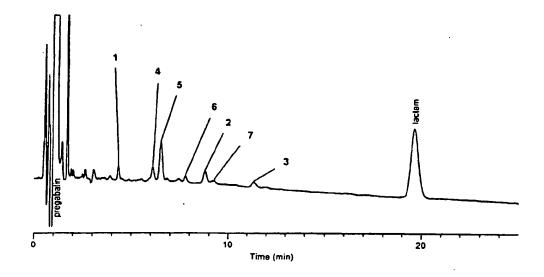


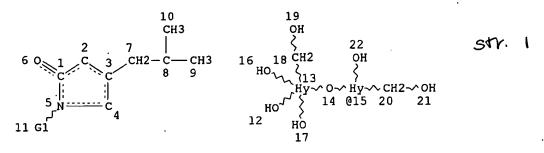
Figure 3. HPLC Chromatogram of Pregabalin 25-mg Capsules Stored at 40°C/75% RH for 6 Months (Modified Method) HPLC Conditions; Column: Phenomenex Luna C18 (2) 100 × 4.6 mm ID, 3 μ; Mobile Phase: 85:15 0.05% Formic Acid:CH₃CN; Flow Rate: 1.5 mL/min; Detection: 210 nm; Temperature: Ambient; Compounds: as defined in Scheme 3

FILE 'CAPLUS' ENTERED AT 12:03:20 ON 18 JAN 2005
L1 0 SEA ABB=ON PLU=ON (4(W)(ISOBUTYL OR (ISO OR I)(W)(BU OR BUTYL)))(S)(4(W)(TRIHYDROXY? OR TRI HYDROXY?))

FILE 'MEDLINE, BIOSIS, EMBASE, WPIDS, CONFSCI, SCISEARCH, JICST-EPLUS, JAPIO' ENTERED AT 12:04:04 ON 18 JAN 2005
L2 0 SEA ABB=ON PLU=ON L1

FILE 'HOME' ENTERED AT 12:06:02 ON 18 JAN 2005

(FILE 'REGISTRY' ENTERED AT 14:46:48 ON 12 JAN 2005) STR



VAR G1=15/23/34 NODE ATTRIBUTES: CONNECT IS X2 RC AT 2 CONNECT IS X2 RC AT 4 CONNECT IS X2 RC AT 33 DEFAULT MLEVEL IS ATOM GGCAT IS SAT AT 13 GGCAT IS SAT AT 15 24 GGCAT IS SAT AT IS SAT ΑT 26 GGCAT DEFAULT ECLEVEL IS LIMITED ECOUNT IS El O AT ECOUNT IS El O AΤ 15 IS E1 O AT ECOUNT 24 ECOUNT IS E1 O AT

GRAPH ATTRIBUTES: RSPEC I

L7

NUMBER OF NODES IS 43

STEREO ATTRIBUTES: NONE

L9 . 12 SEA FILE=REGISTRY SSS FUL L7

100.0% PROCESSED 14218 ITERATIONS

(FILE 'CAPLUS' ENTERED AT 15:07:11 ON 12 JAN 2005)

L10 2 S L9

SEARCH TIME: .00.00.02

L10 ANSWER 1 OF 2 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER: 2002:777769 CAPLUS

> Searcher : Shears 571-272-2528

12 ANSWERS

DOCUMENT NUMBER:

137:284373

Pregabalin-lactose conjugates for pharmaceuticals TITLE: Hurley, Timothy Robert; Lovdahl, Michael James; INVENTOR(S):

Tobias, Brian

PATENT ASSIGNEE(S):

Warner-Lambert Company, USA

SOURCE:

PCT Int. Appl., 36 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

	PA	CENT :	NO.					DATE							. OV		. D.	ATE	
	WO	2002	0787					2002	1010)2-:		 7		2	0020	225
		2002						2003	1009										
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	WO	2002	0787	47		C2		2003	1231										
		W:	AE,	AG,	AL,	AM,	AT,	ΑU,	AZ,	BA,	BB	3, E	3G,	BR,	BY,	BZ,	CA,	CH,	CN,
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								TM,											
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AΒ Compns. containing pregabalin-lactose conjugates are useful for the treatment

of of central nervous system disorders or diseases including seizure disorders, pain, depression, anxiety, sleep disorders, consumptive disorders, psychosis, tardive dyskinesia, Huntington's disease, or Parkinson's disease in humans. Pregabalin and lactose were dissolved in water and the solution was then heated at 90°. The resulting solid was then redissolved in approx. iso-PrOH by sonicating and heating. The product was subjected to reversed-phase preparative chromatog. to give $(S)-1-\{3,4-dihydroxy-6-hydroxymethyl-5-(3,4,5-trihydroxymethyl-6$ hydroxymethyltetrahydropyran-2-yloxy) tetrahydropyran-2-yl]-4isobutylpyrrolidin-2-one (I). Tablets contained I 25, lactose 50, corn starch (for mix) 10, corn starch (paste) 10, and Mg stearate 5 mg.

IT 466678-44-4P 466678-45-5P 466678-46-6P 466678-47-7P 466678-50-2P

RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological

Searcher :

Shears

571-272-2528

Absolute stereochemistry.

RN 466678-45-5 CAPLUS CN β -D-Fructofuranose, 1-deoxy-4-O- β -D-galactopyranosyl-1-[(45)-4-(2-methylpropyl)-2-oxo-1-pyrrolidinyl]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 466678-46-6 CAPLUS
CN β-D-Fructopyranose, 1-deoxy-4-O-β-D-galactopyranosyl-1-[(4S)-4-(2-methylpropyl)-2-oxo-1-pyrrolidinyl]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 466678-47-7 CAPLUS
CN α-D-Tagatopyranose, 1-deoxy-1-[(4S)-4-(2-methylpropyl)-2-oxo-1pyrrolidinyl]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 466678-50-2 CAPLUS
CN β-D-Tagatopyranose, 1-deoxy-1-[(4S)-4-(2-methylpropyl)-2-oxo-1pyrrolidinyl]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

IT 466678-49-9

> RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (pregabalin lactose conjugates for pharmaceuticals)

RN 466678-49-9 CAPLUS

B-D-Fructopyranose, 1-deoxy-1-[(4S)-4-(2-methylpropyl)-2-oxo-1-CN pyrrolidinyl) - (9CI) (CA INDEX NAME)

Absolute stereochemistry.

L10 ANSWER 2 OF 2 CAPLUS COPYRIGHT 2005 ACS on STN

2002:402961 CAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 138:243397

TITLE: Synthesis and characterization of pregabalin lactose

conjugate degradation products

Lovdahl, Michael J.; Hurley, Timothy R.; Tobias, AUTHOR(S):

Brian; Priebe, Stephen R.

CORPORATE SOURCE: Analytical Development Department, Pfizer Global

Research and Development, Ann Arbor, MI, 48105, USA

Journal of Pharmaceutical and Biomedical Analysis (2002), 28(5), 917-924
CODEN: JPBADA; ISSN: 0731-7085 SOURCE:

PUBLISHER: Elsevier Science B.V.

DOCUMENT TYPE: Journal LANGUAGE: English

Seven degradation products observed in formulated pregabalin have been characterized. These compds. result from Maillard reactions and Amadori rearrangements. Heating pregabalin in the presence of lactose formed significant quantities of these degradation products. The seven compds.

corresponding to the observed degradation products were isolated by

preparative liquid chromatog. The synthesis, isolation, and spectral characterization of the degradation products are detailed.

501665-97-0, PD 224377 501666-22-4, PD 310806

501666-23-5, PD 312237 501666-24-6, PD 312236

501666-25-7, PD 310886

RL: FMU (Formation, unclassified); PRP (Properties); FORM (Formation, nonpreparative)

(synthesis and characterization of pregabalin lactose conjugate degradation

products)

RN 501665-97-0 CAPLUS

CN \(\alpha - D - Tagatopyranose, \) 1 - deoxy-1 - [(4R) - 4 - (2 - methylpropyl) - 2 - 0x0 - 1 - pyrrolidinyl] - (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 501666-22-4 CAPLUS

CN β -D-Tagatopyranose, 1-deoxy-1-[(4R)-4-(2-methylpropyl)-2-oxo-1-pyrrolidinyl]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 501666-23-5 CAPLUS

CN β-D-Fructofuranose, 1-deoxy-4-O-β-D-galactopyranosyl-1-[(4R)-4-(2-methylpropyl)-2-oxo-1-pyrrolidinyl)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Searcher :

Shears

571-272-2528

RN 501666-24-6 CAPLUS

CN β -D-Fructopyranose, 1-deoxy-4-O- β -D-galactopyranosyl-1-[(4R)-4-(2-methylpropyl)-2-oxo-1-pyrrolidinyl]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 501666-25-7 CAPLUS

CN β-D-Fructopyranose, 1-deoxy-1-{(4R)-4-(2-methylpropyl)-2-oxo-1-pyrrolidinyl]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

IT 501665-88-9, PD 224378

RL: FMU (Formation, unclassified); PRP (Properties); RCT (Reactant); FORM (Formation, nonpreparative); RACT (Reactant or reagent) (synthesis and characterization of pregabalin lactose conjugate

degradation

products)

RN 501665-88-9 CAPLUS

CN 2-Pyrrolidinone, $1-(4-0-\beta-D-galactopyranosyl-\beta-D-glucopyranosyl)-4-(2-methylpropyl)-, (4R)- (9CI) (CA INDEX NAME)$

Absolute stereochemistry.

REFERENCE COUNT:

THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

FILE 'CAOLD' ENTERED AT 15:07:57 ON 12 JAN 2005 L11 0 S L9

FILE 'USPATFULL' ENTERED AT 15:08:03 ON 12 JAN 2005 L12 1 S L9

L12 ANSWER 1 OF 1 USPATFULL on STN

ACCESSION NUMBER: 2002:330258 USPATFULL

TITLE: Pregabalin lactose conjugates

INVENTOR(S): Hurley, Timothy Robert, Ann Arbor, MI, UNITED STATES

Lovdahl, Michael James, Ann Arbor, MI, UNITED STATES

Tobias, Brian, Ann Arbor, MI, UNITED STATES

	NUMBER	KIND	DATE		alphi
PATENT INFORMATION: APPLICATION INFO.:	US 2002187941 US 2002-58903	A1 A1	20021212 20020128	(10)	Yms

NUMBER DATE

PRIORITY INFORMATION: US 2001-280176P 20010330 (60)

DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: David R. Kurlandsky, Warner-Lambert Company, 2800

Plymouth Road, Ann Arbor, MI, 48105

NUMBER OF CLAIMS: 7
EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 3 Drawing Page(s)

LINE COUNT:

891

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB In accordance with the present invention, there is provided pregabalin lactose conjugate compounds.

Also provided as part of the present invention is a novel method of central nervous system disorders or diseases including seizure disorders, pain, depression, anxiety, sleep disorders, consumptive disorders, psychosis, tardive dyskinesia, Huntington's disease, or Parkinson's disease in a subject by administering to the subject a pharmaceutically effective amount of a pregabalin lactose conjugate.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

(FILE 'MEDLINE, BIOSIS, EMBASE' ENTERED AT 15:08:26 ON 12 JAN 2005) L13 0 S L9

(FILE 'MARPAT' ENTERED AT 15:08:41 ON 12 JAN 2005)

VAR G1=15/23/34 NODE ATTRIBUTES: CONNECT IS X2 RC AT CONNECT IS X2 RC AT CONNECT IS X2 RC AT 33 DEFAULT MLEVEL IS ATOM MLEVEL IS CLASS AT 13 15 24 26 GGCAT IS SAT AT 13 GGCAT IS SAT AT GGCAT IS SAT AT GGCAT IS SAT AT DEFAULT ECLEVEL IS LIMITED ECOUNT IS E1 O AT 13

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GRAPH ATTRIBUTES:

RSPEC I

NUMBER OF NODES IS 43

STEREO ATTRIBUTES: NONE

ATTRIBUTES SPECIFIED AT SEARCH-TIME:

ECLEVEL IS LIM ON ALL NODES ALL RING(S) ARE ISOLATED

L16

O SEA FILE=MARPAT SSS FUL L14 (MODIFIED ATTRIBUTES)

100.0% PROCESSED 5780 ITERATIONS

O ANSWERS

SEARCH TIME: 00.00.05

FILE 'MARPATPREV' ENTERED AT 15:09:51 ON 12 JAN 2005 L14 STR

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CONNECT IS X2 RC AT 4

CONNECT IS X2 RC AT 33 DEFAULT MLEVEL IS ATOM

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GGCAT IS SAT AT 13

AT 15 IS SAT GGCAT

GGCAT IS SAT AT 24

IS SAT AT 26 GGCAT

DEFAULT ECLEVEL IS LIMITED

ECOUNT IS El O AT 13

571-272-2528 Searcher : Shears

ECOUNT IS ELO AT 15 ECOUNT IS El O AT 24 ECOUNT IS E1 O AT 26 GRAPH ATTRIBUTES: RSPEC I NUMBER OF NODES IS 43 STEREO ATTRIBUTES: NONE ATTRIBUTES SPECIFIED AT SEARCH-TIME: ECLEVEL IS LIM ON ALL NODES ALL RING(S) ARE ISOLATED O SEA FILE=MARPATPREV SSS FUL L14 (MODIFIED ATTRIBUTES) L17 0 ANSWERS 100.0% PROCESSED 12 ITERATIONS SEARCH TIME: 00.00.01 (FILE 'REGISTRY' ENTERED AT 15:13:22 ON 12 JAN 2005) 0 S ?"ISOBUTYL-1-(2,3,4,5-TETRAHYDROXY"?/CNS L25 - Named compds O S ?"ISOBUTYL-1-(2,3,5-TRIHYDROXY"?/CNS L27 0 S ?"ISOBUTYL-1-(2,3,4-TRIHYDROXY"?/CNS L28 0 S ?"DIHYDROXY-6-HYDROXYMETHYL-5-(3,4,5-TRIHYDROXY"?/CNS L31 0 S ?"DIHYDROXY-5-HYDROXYMETHYL-4-(3,4,5-TRIHYDROXY"?/CNS L32 (FILE 'CAPLUS' ENTERED AT 15:20:41 ON 12 JAN 2005) L33 1381 SEA FILE=CAPLUS ABB=ON PLU=ON (DIHYDROXY OR DI HYDROXY) (S) (HY DROXYMETHYL OR HYDROXY(W) (ME OR METHYL)) 128 SEA FILE=CAPLUS ABB=ON PLU=ON L33(S) (TETRAHYDRO? OR TETRA L34 HYDRO?) 1 SEA FILE=CAPLUS ABB=ON PLU=ON L34(S) (ISOBUTYL? OR (ISO OR L35 I) (W) (BU OR BUTYL?)) 692 SEA FILE=CAPLUS ABB=ON PLU=ON 4(W) (ISOBUTYL OR (ISO OR L36 I) (W) (BUTYL OR BU)) O SEA FILE=CAPLUS ABB=ON PLU=ON L36(S)(4(W)5(W)(TETRAHYDROXY? L37 OR TETRA HYDROXY?)) 692 SEA FILE=CAPLUS ABB=ON PLU=ON 4(W) (ISOBUTYL OR (ISO OR L36 I) (W) (BUTYL OR BU)) O SEA FILE=CAPLUS ABB=ON PLU=ON L36(S)(5(W)(TRIHYDROXY? OR TRI L38 HYDROXY?)) 0 L35 NOT L10 L39 (FILE 'MEDLINE, BIOSIS, EMBASE, WPIDS, CONFSCI, SCISEARCH, JICST-EPLUS, JAPIO' ENTERED AT 15:25:06 ON 12 JAN 2005) L40 1 S L35

Shears

571-272-2528

Searcher :

L41 0 S L37 L42 0 S L38

L40 ANSWER 1 OF 1 WPIDS COPYRIGHT 2005 THE THOMSON CORP on STN

ACCESSION NUMBER: 2002-241176 [29] WPIDS

DOC. NO. CPI: C2002-072446

TITLE: New beta-hydroxyamide compound useful in powder coating

compositions as crosslinkers and/or curing agents.

DERWENT CLASS: A25 A60 E16 G02
INVENTOR(S): MANEA, M; PETERSSON, C

PATENT ASSIGNEE(S): (PEST) PERSTORP SPECIALTY CHEM AB; (MANE-I) MANEA M;

(PETE-I) PETERSSON C

COUNTRY COUNT: 95

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 2001098257 Al 20011227 (200229)* EN 23

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ

NL OA PT SD SE SL SZ TR TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC

LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE

SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW

AU 2001074750 A 20020102 (200230)

EP 1292567 A1 20030319 (200322) EN

R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT

RO SE SI TR

US 2003195373 A1 20031016 (200369)

JP 2004501133 W 20040115 (200410) 42

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 2001098257	A1	WO 2001-SE1359	20010615
AU 2001074750	A	AU 2001-74750	20010615
EP 1292567	A1	EP 2001-941393	20010615
		WO 2001-SE1359	20010615
US 2003195373	A1	WO 2001-SE1359	20010615
		US 2003-311295	20030204
JP 2004501133	W	WO 2001-SE1359	20010615
		JP 2002-504213	20010615

FILING DETAILS:

PATENT NO	KIND	PATENT NO
AU 2001074750	A Based on	WO 2001098257
EP 1292567	Al Based on	WO 2001098257
JP 2004501133	W Based on	WO 2001098257

PRIORITY APPLN. INFO: SE 2000-2268 20000619

AN 2002-241176 [29] WPIDS

AB WO 200198257 A UPAB: 20020508

NOVELTY - A beta -hydroxyamide compound (I) is new.

DETAILED DESCRIPTION - A beta -hydroxyamide compound of formula (I) is new.

R1 = alkyl, alkoxyalkyl, hydroxyalkyl or hydroxyalkoxyalkyl;

R2 = alkyl, aryl, alkylaryl or arylalkyl;

R3 = N-alkyl or N-cycloalkyl having at least one OH in beta
-position;

m, n = 1 or more.

An INDEPENDENT CLAIM is also included for a process for the synthesis of (\mathbf{I}) .

USE - In powder coating compositions as chemical intermediates, crosslinkers and/or curing agents.

ADVANTAGE - (I) provides a coating that can be formulated as a solvent-borne or water-borne system, and solvent-borne coatings can be formulated as high-solid systems. It allows crosslinking temperature to be moderate-to-high in the range of 150-200 deg. C, and reduces the compatibility problems due to di, tri or polyhydric core compounds. The coating obtained is clear and has improved flexibility and hardness. Dwg.0/0

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